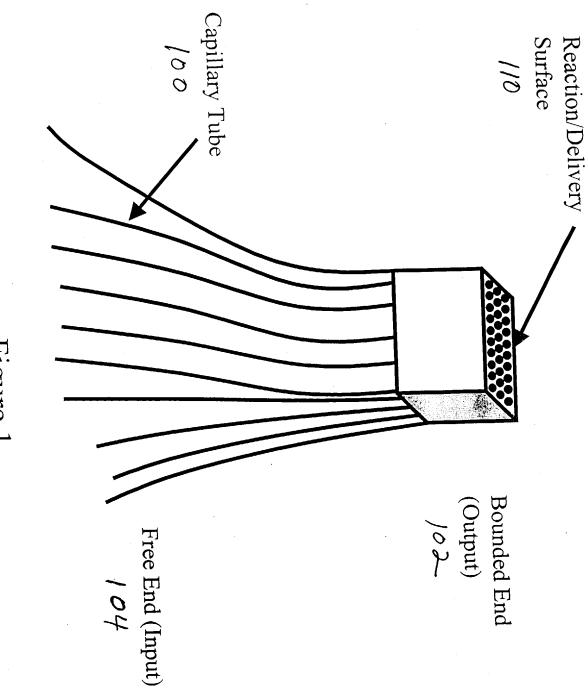
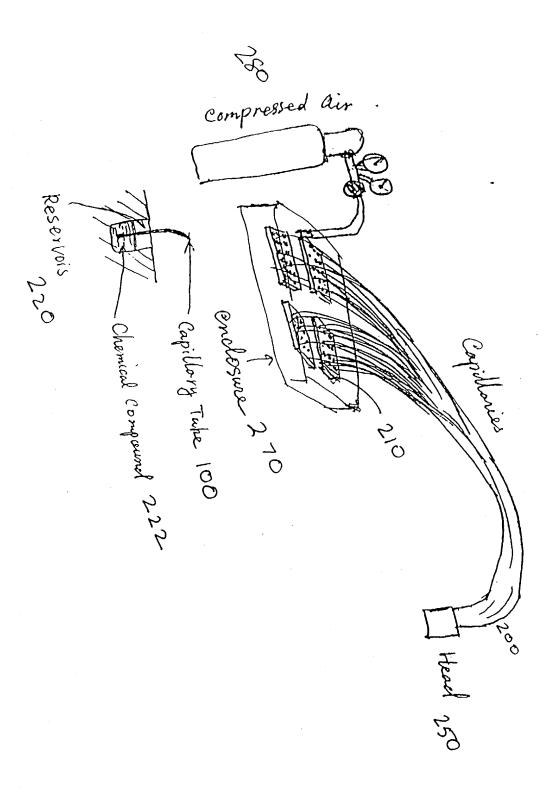
Title: THOD AND APPARATUS BASED ON BUNDLED CAPPARATES FOR THROUGHPUT SCREENING Inventor. Jianming XIAO et al. Application No.: To Be Assigned Docket No.: 473532000620

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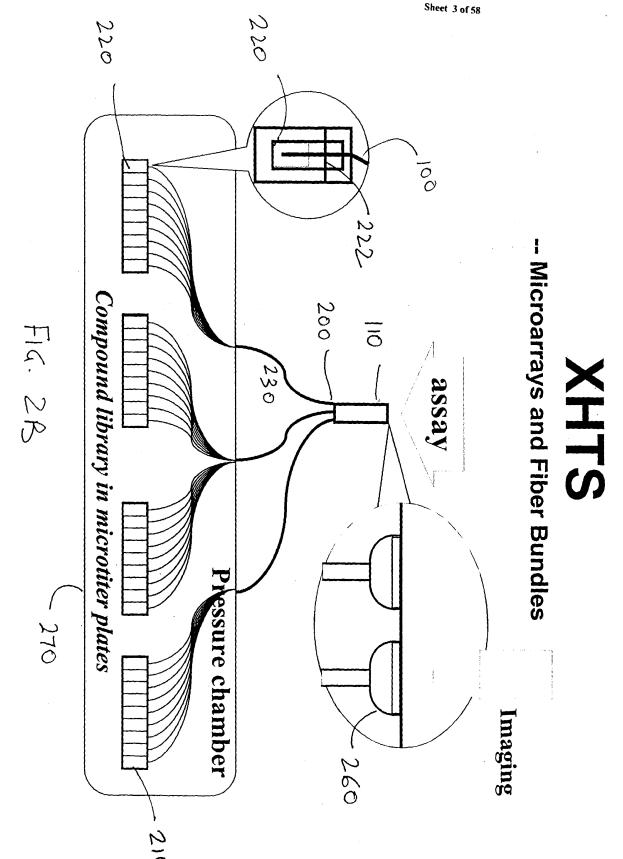


Title: MET AND APPARATUS BASED ON BUNDLED CAPILLA FOR HIGH JUGHPUT SCREENING Inventor: Jianming XIAO et al. Application No.: To Be Assigned Docket No.: 473532000620

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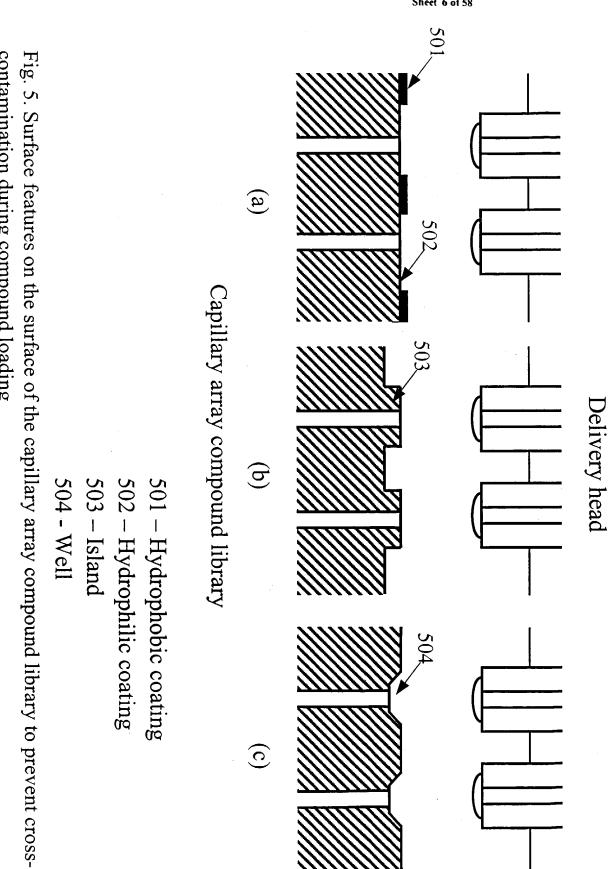


Title: METHOD AND APPARATUS BASED ON TOLED CAPILLARIES FOR HIGH THROUGHPUT SCREENING Inventor: Jianming XIAO et al.
Application No.: 473532000620 Sheet 5 of 58 B Fig. 4. Fabrication of delivery head using a guide plate 0-(

ND APPARATUS BASED ON BUNDLED CAPILLAR IGHPUT SCREENING Title: METHO FOR HIGH 7

Inventor: Jianming XIAO et al. Application No.: To Be Assigned Docket No.: 473532000620

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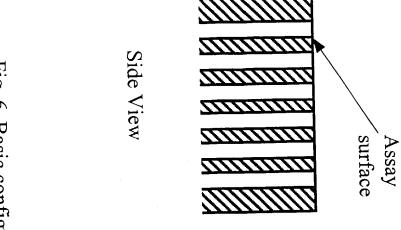


contamination during compound loading

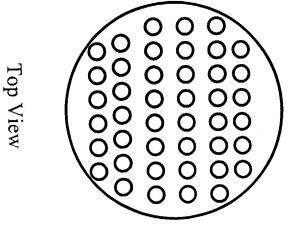
Title: ME TOD AND APPARATUS BASED ON BUNDLED CAPILL SIES FOR HIGH ROUGHPUT SCREENING

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portable compound library Fig. 6. Basic configuration of capillary array substrate for the



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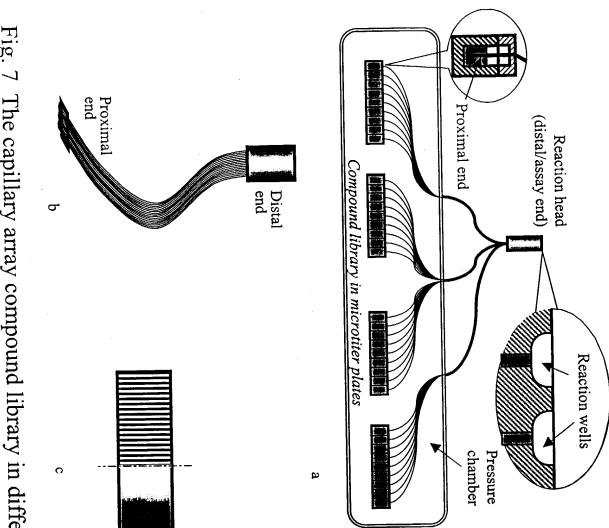


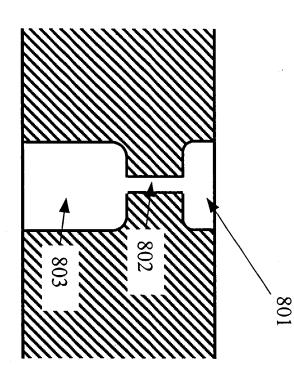
Fig. 7 The capillary array compound library in different formats

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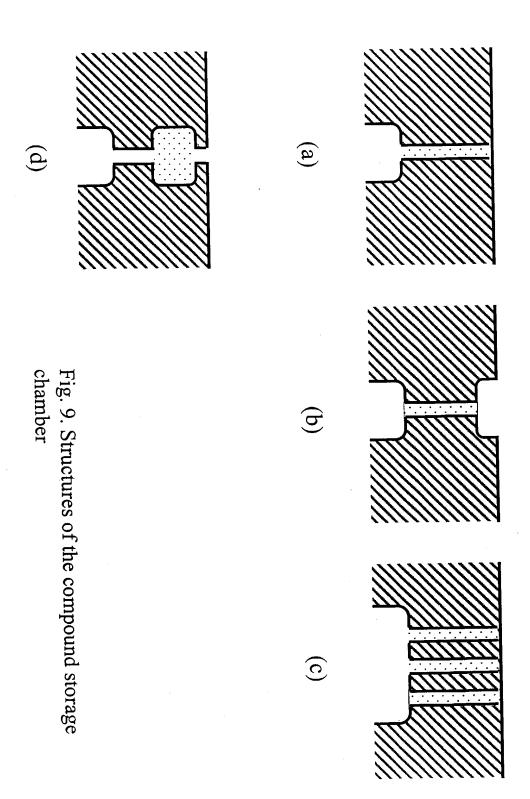
Fig. 8. Internal structure of a through hole in capillary array compound library



801 – Mixing/reaction well 802 – Flow regulator for reagent metering 803 – Compound reservoir

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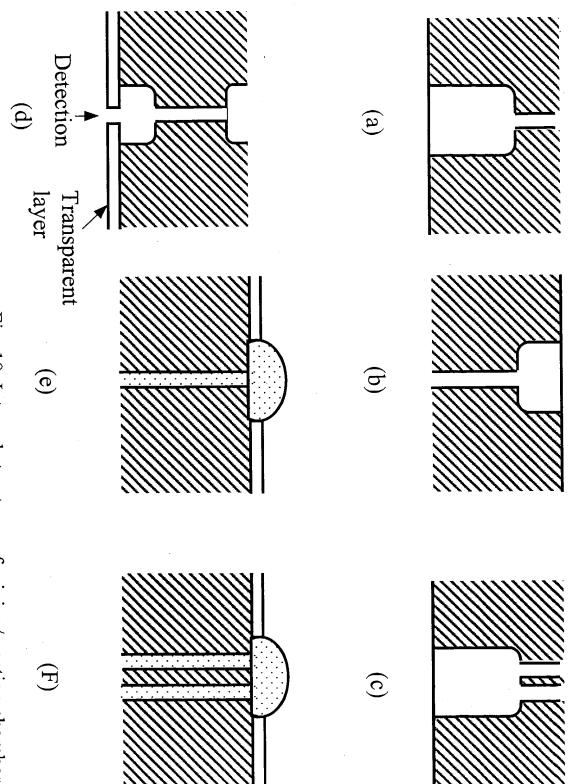


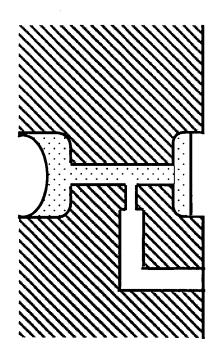
Fig. 10. Internal structures of mixing/reaction chamber

AND APPARATUS BASED ON BUNDLED CAPILLARIES OUGHPUT SCREENING Title: MET FOR HIG Inventor: Jianming XIAO et al. Application No.: To Be Assigned Docket No.: 473532000620 Sheet 12 of 58 (a) <u>C</u> **(b)** tension patch Fig. 11. Volume metering by surface 1101 – Hydrophobic coating1102 – Hydrophilic coating

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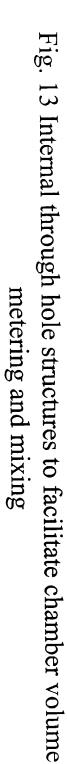
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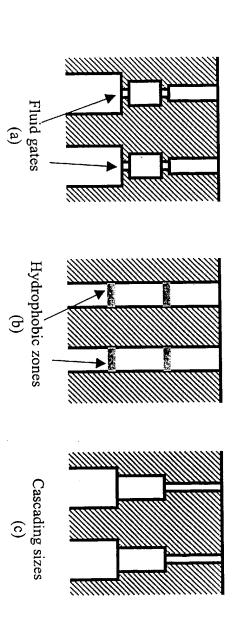
Fig. 12. Fluid regulator with side air tunnel



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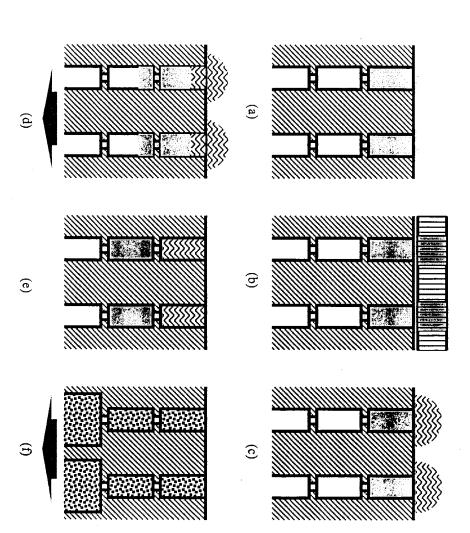




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g. 14 Process of metering multiple reagents using interconnected chambers

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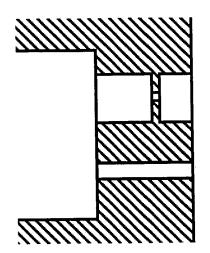
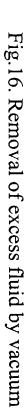
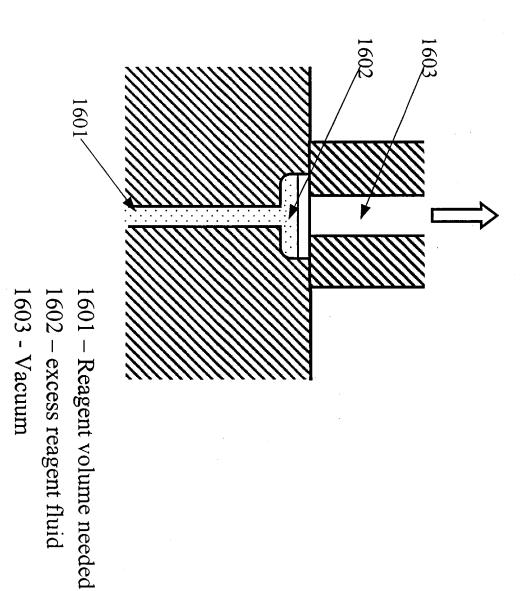


Fig. 15 Special through hole structure where multiple chambers links to a chamber in parallel

Inventor: Jianming XIAO et al. Application No.: To Be Assigned Docket No.: 473532000620

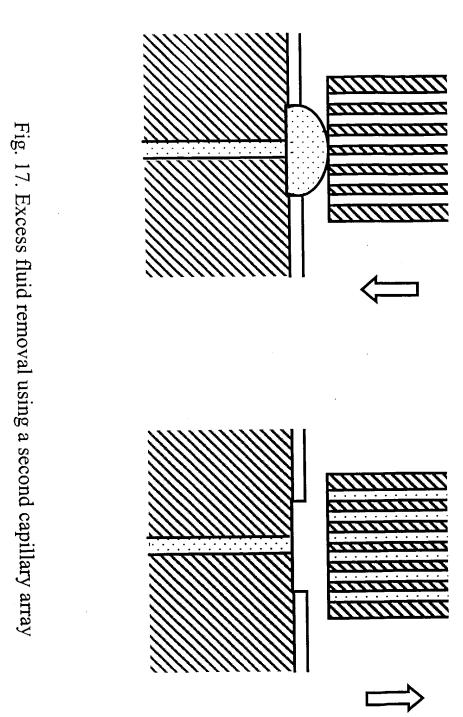
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Title: MET OD AND APPARATUS BASED ON BUNDLED CAPUS FOR HEAD AROUGHPUT SCREENING Inventor: Jianming XIAO et al. Application No.: To Be Assigned Docket No.: 473532000620 PIES

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Title: ME OD AND APPARATUS BASED ON BUNDLED CAPIL RIES FOR HOUGHPUT SCREENING

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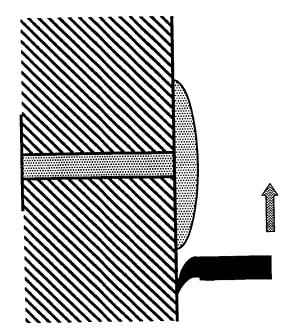
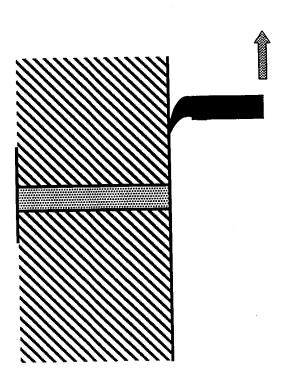


Fig. 18. Excess Fluid Removal by Wiping

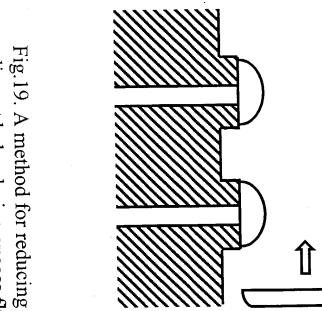


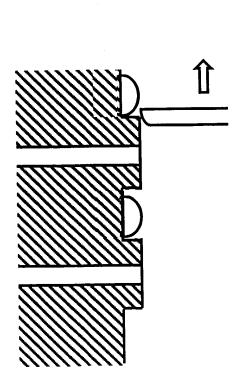
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Title: MFT OD AND APPARATUS BASED ON BUNDLED CAPIL FOR HI IROUGHPUT SCREENING Inventor: Jianming XIAO et al. Application No.: To Be Assigned Docket No.: 473532000620

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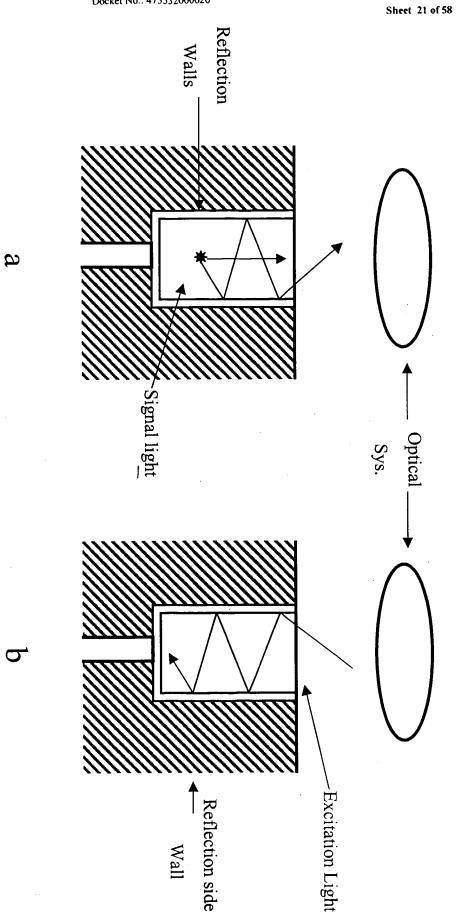




adjacent holes during excess fluid removal Fig. 19. A method for reducing cross-contamination between Title: MET 'OD AND APPARATUS BASED ON BUNDLED CAPILL 'ZIES FOR HE HOUGHPUT SCREENING Inventor: Jianming XIAO et al.

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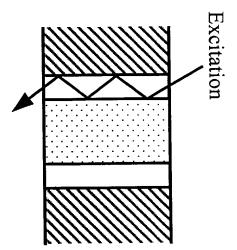
METHOD AND APPARATUS BASED ON BUNDLED CAPILLARIES

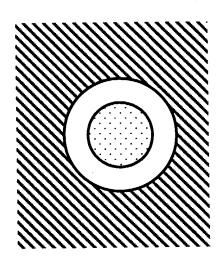
GH THROUGHPUT SCREENING
Inventor: Jianming XIAO et al.

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Title: METHES AND APPARATUS BASED ON BUNDLED CAPILLAPIES OUGHPUT SCREENING FOR HIGH

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R=-04,-C/4,-C/4-

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FIG 22B

COOH

Title: METHOD AND APPARATUS BASED ON BUNDLED CAPILLAPIES FOR HIG OUGHPUT SCREENING

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FIG. 22C

Figure 2.3 Process for fabrication using a negative mask

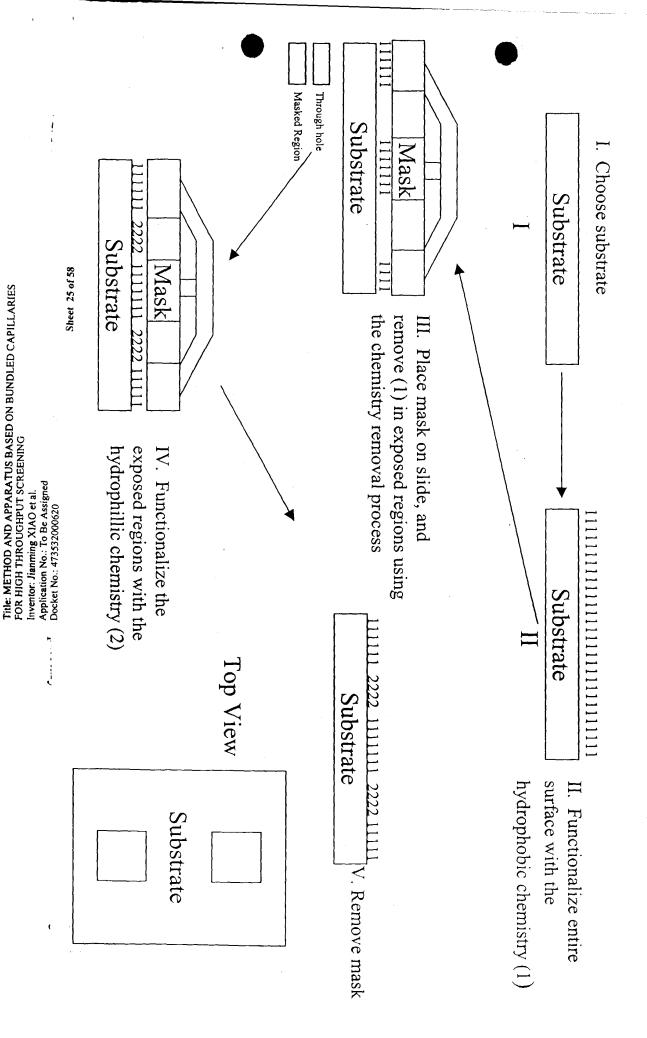


Figure 24 Process for the fabrication using positive mask

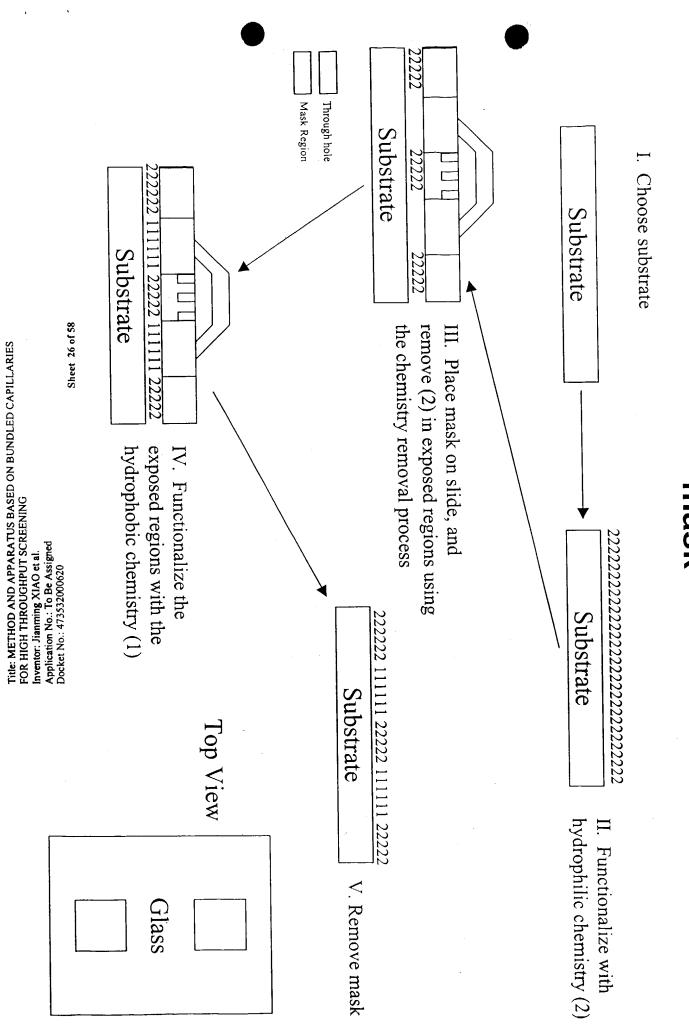
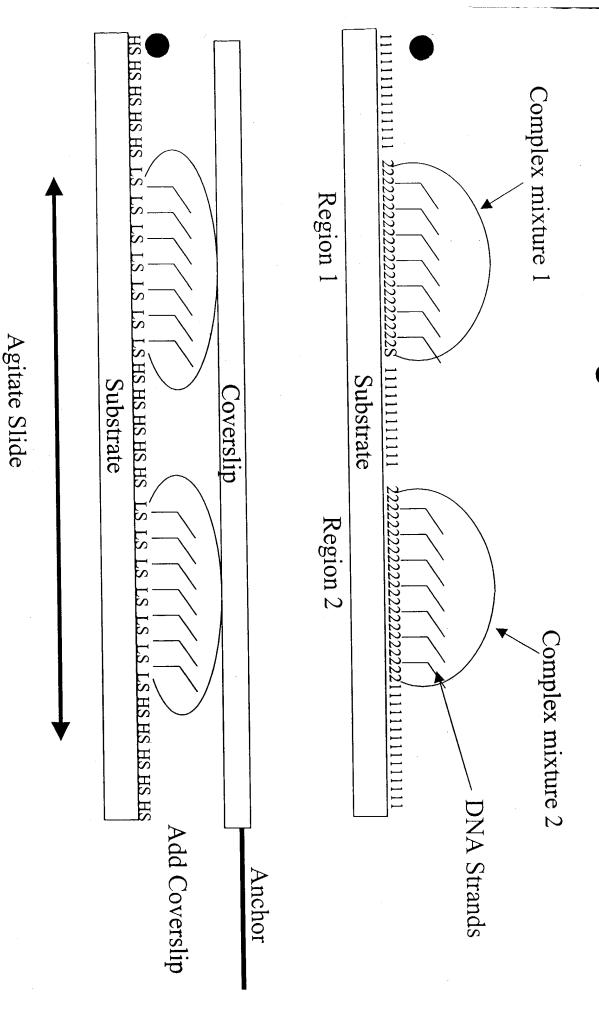


Figure 25 Chamber use

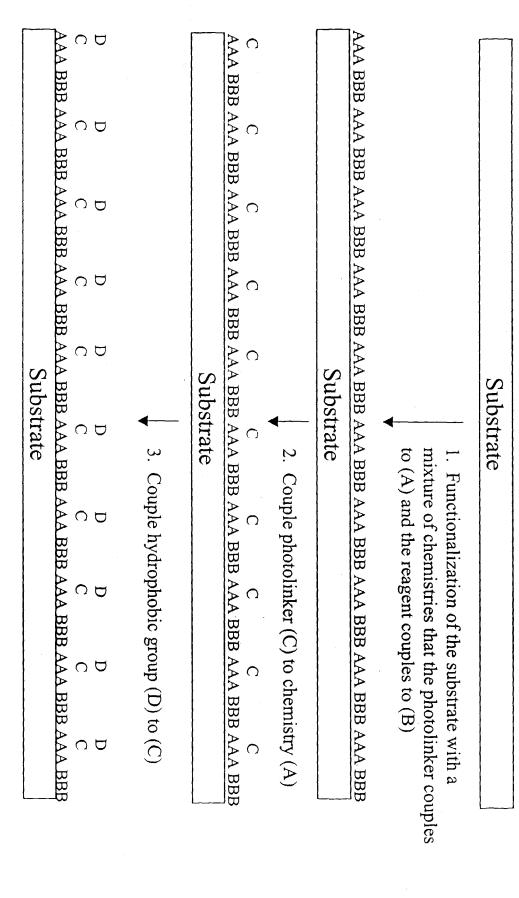


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Title: METHOD AND APPARATUS BASED ON BUNDLED CAPILLARIES FOR HIGH THROUGHPUT SCREENING Inventor: Jianming XIAO et al. Application No.: To Be Assigned Docket No.: 473532000620

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Figure 26A Surface Tension Patterning: On-capillary Fiber optic based patterning



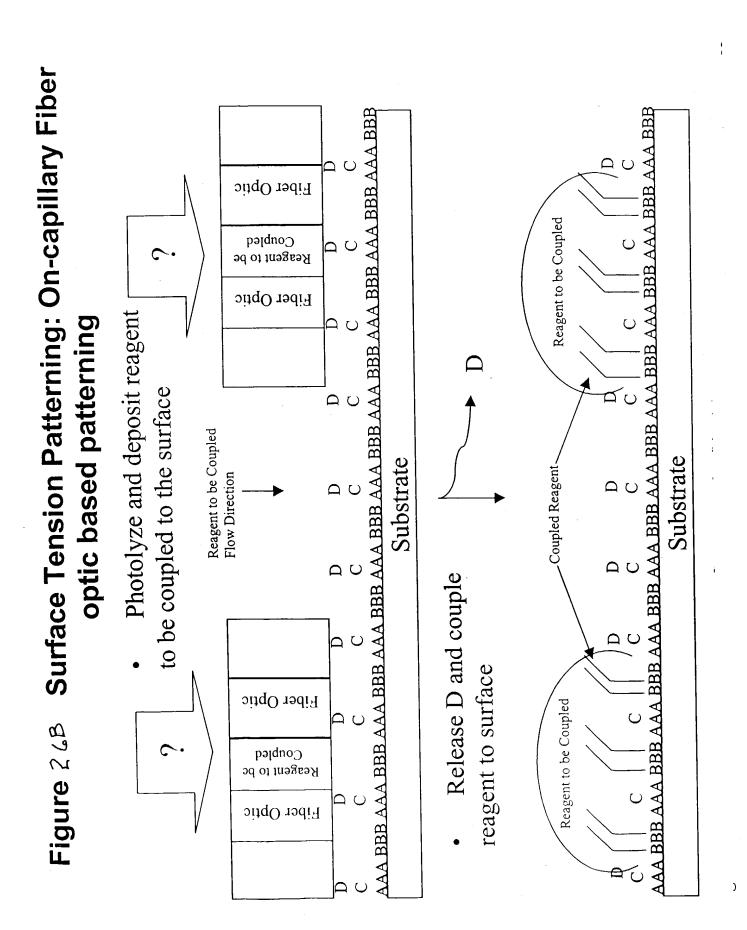
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Title: METHOD AND APPARATUS BASED ON BUNDLED CAPILLARIES FOR HIGH THROUGHPUT SCREENING Inventor: Jianming XIAO et al.
Application No.: To Be Assigned Docket No.: 473332000620

Title: METL ND APPARATUS BASED ON BUNDLED CAPILLARIES FOR HIGH AUGHPUT SCREENING Lawrenter: Lianning XIAO et al.

Inventor: Jianming XIAO et al. Application No.: To Be Assigned Docket No.: 473532000620

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Glass Fiber Optic

Fiber Optic Glass Glass Fiber Optic

Fiber Optic Glass

Glass Fiber Optic

Fiber Optic Glass Glass Fiber Optic

Fiber Optic Glass

Coat Surface with a Hydrophobic Reagent

Figure 27A Volume Metering using Surface **Tension Features**

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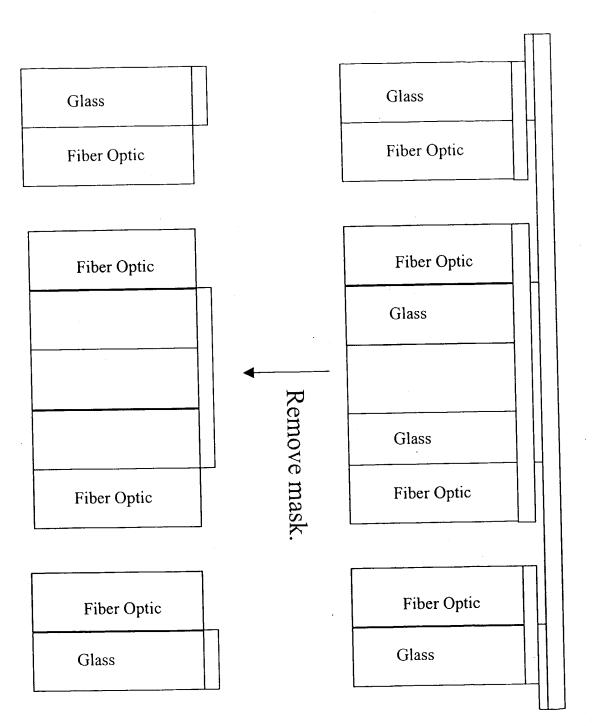


Figure 276 Volume Metering using Surface Tension Features

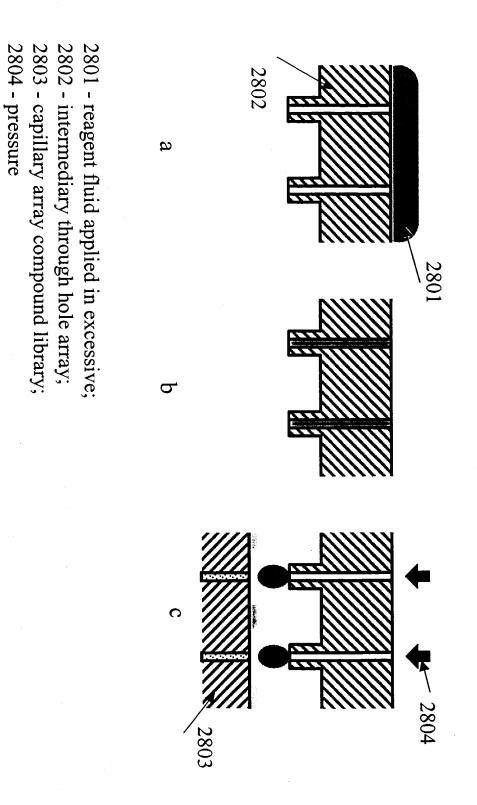
Place a Mask on to the Surface and Expose the Surface to the Chemistry Removal Process

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Fig. 28 Reagent pre-metering using an intermediary through-hole array

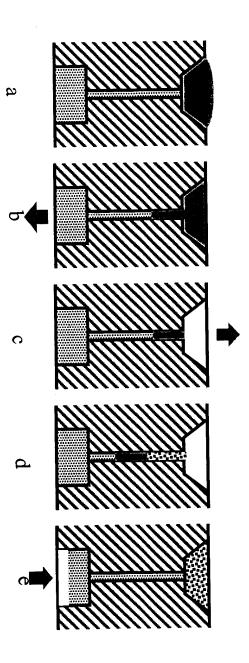


Title: METHO TO APPARATUS BASED ON BUNDLED CAPILLARIES FOR HIGH TO THE TOTAL SCREENING

Inventor: Jianming XIAO et al.
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Title: METHS D AND APPARATUS BASED ON BUNDLED CAPILLARUS FOR HIGH OUGHPUT SCREENING Inventor: Jianning XIAO et al.

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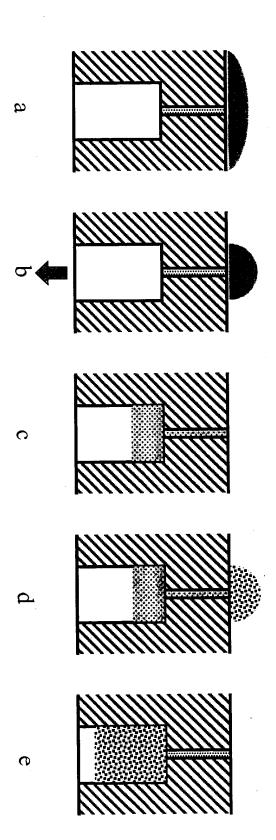


Fig. 30 Metering with hydrophilic patch and mixing

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FOR HIGH
INVENTOR JAMES AND SCREENING
Inventor: Jianning XIAO et al.
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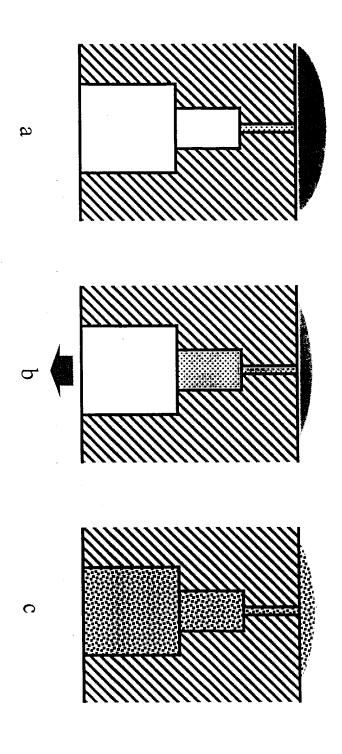


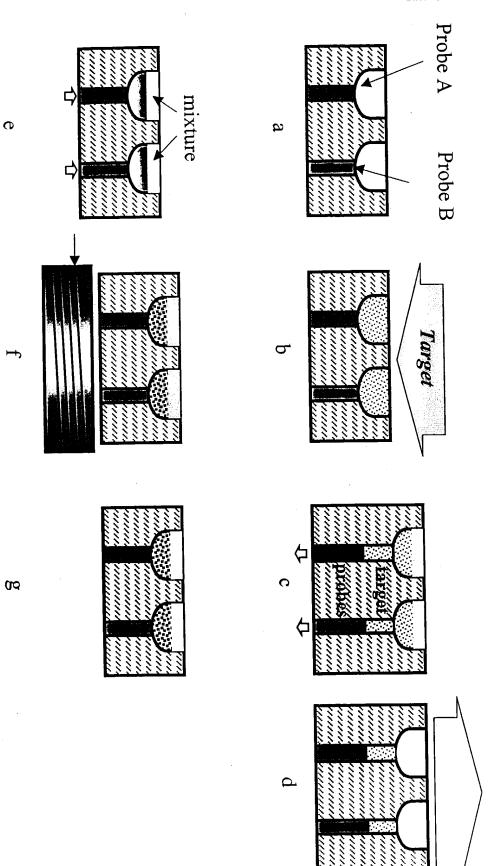
Fig. 31 Mixing and metering with interconnected chambers

Title: MET 4OD AND APPARATUS BASED ON BUNDLED CAPIL! ARIES FOR HROUGHPUT SCREENING

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Fig. 32 Heterogeneous Assay



THROUGHPUT SCREENING
Invented Jianming XIAO et al.
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Sheet 37 of 58 Oxidation of antibodies vicinal diol group to its aldehyde

Antibody Immobilization via the Carbohydrate

Moiety

2. Conjugation of maleimide moiety with antibody

DMSO

3. Immobilization of the modified antibody to the surface.

immobilization
$$0$$
 $-S$
 0
 0
 0
 0
 0
 0
 0
 0

O-NH-N-

FIG. 33A

lmmobilization via Amine Goups

. Hydrosilylation of (3-mercaptopropyl)triethoxysilane on the surface of fiber

$$-OH + (OEt)_3-Si$$
 SH toluene $-O-Si$ SH

2. Formation of a thioether bond

3. Attachment of fiber to antibody

Antibody Immobilization via Streptavidin

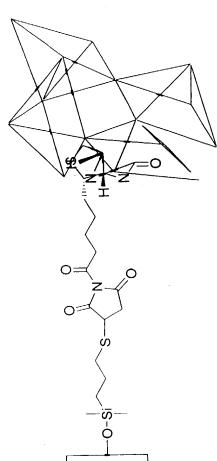
1. Label antibody with biotin

2. Modification of fiber surface with biotin maleimide

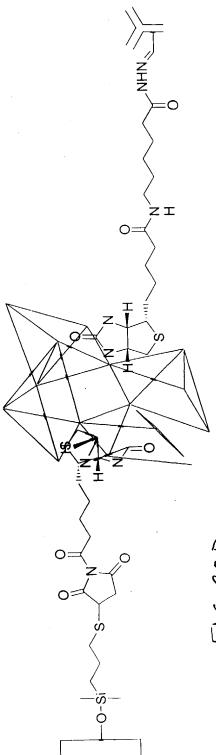
SH H O O
$$\frac{1}{2}$$
 FIG 33 C

Antibody Immobilization via Streptavidin

3. Conjugate Streptavidin to the surface



4. Conjugate Biotin Anitbody to the surface



F16.33D

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Formation of thiazolidine

1. Surface attachment and formation of the linker
$$-0-\frac{1}{2}$$
 $-0-\frac{1}{2}$ $-0-\frac{1}{2}$ $-0-\frac{1}{2}$

IZ

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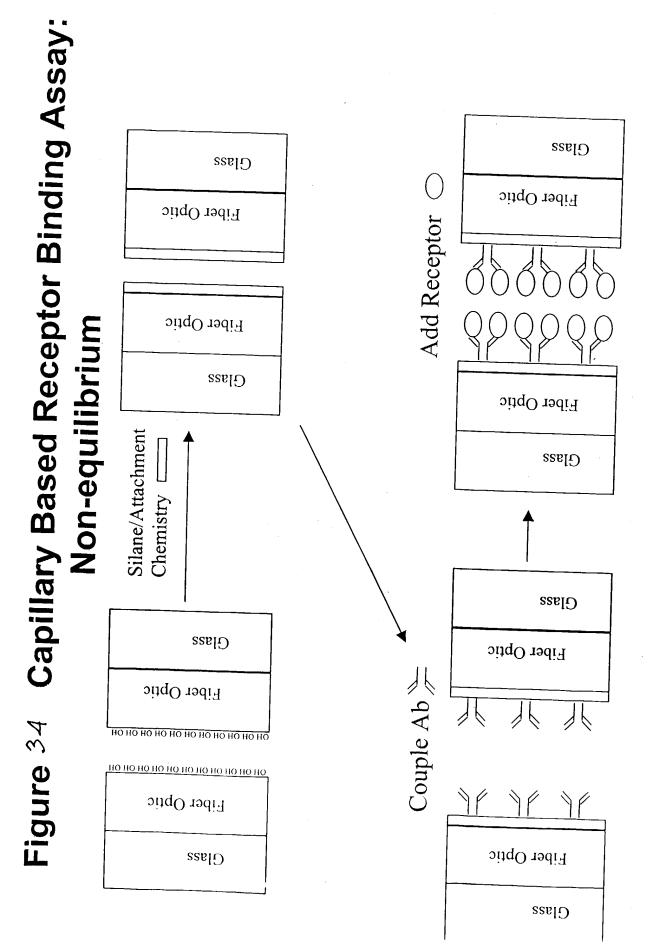
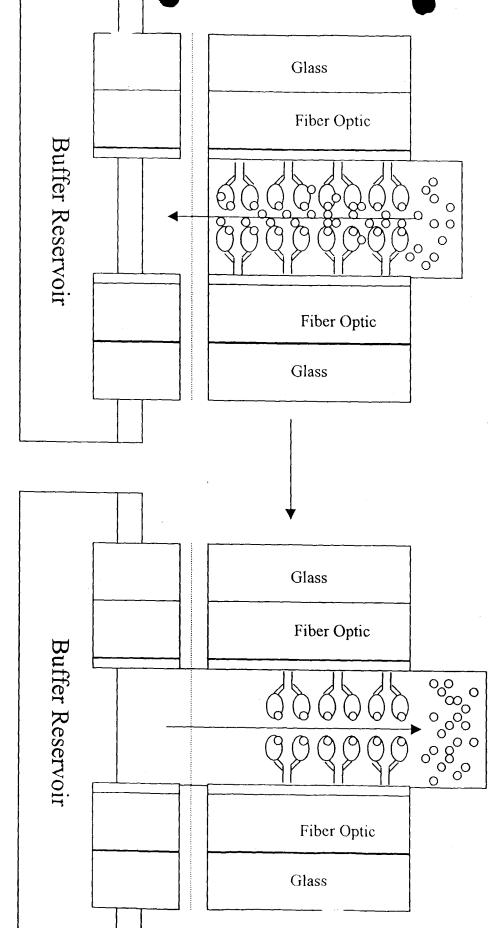


Figure 34 (cont. 1). Capillary Based Receptor Binding Assay: Non-equilibrium

Add saturating ligand

Wash unbound ligand and calculate total bound using fiber optic base detection

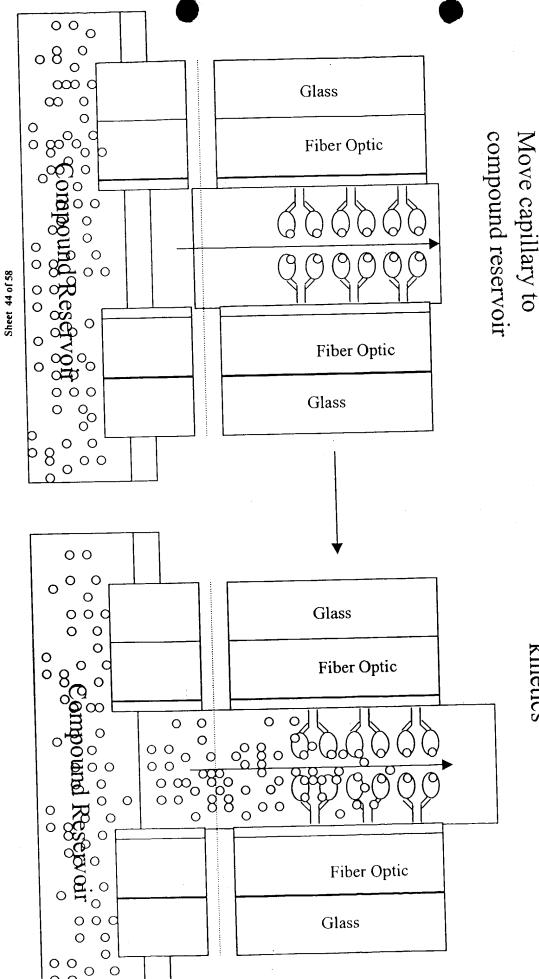


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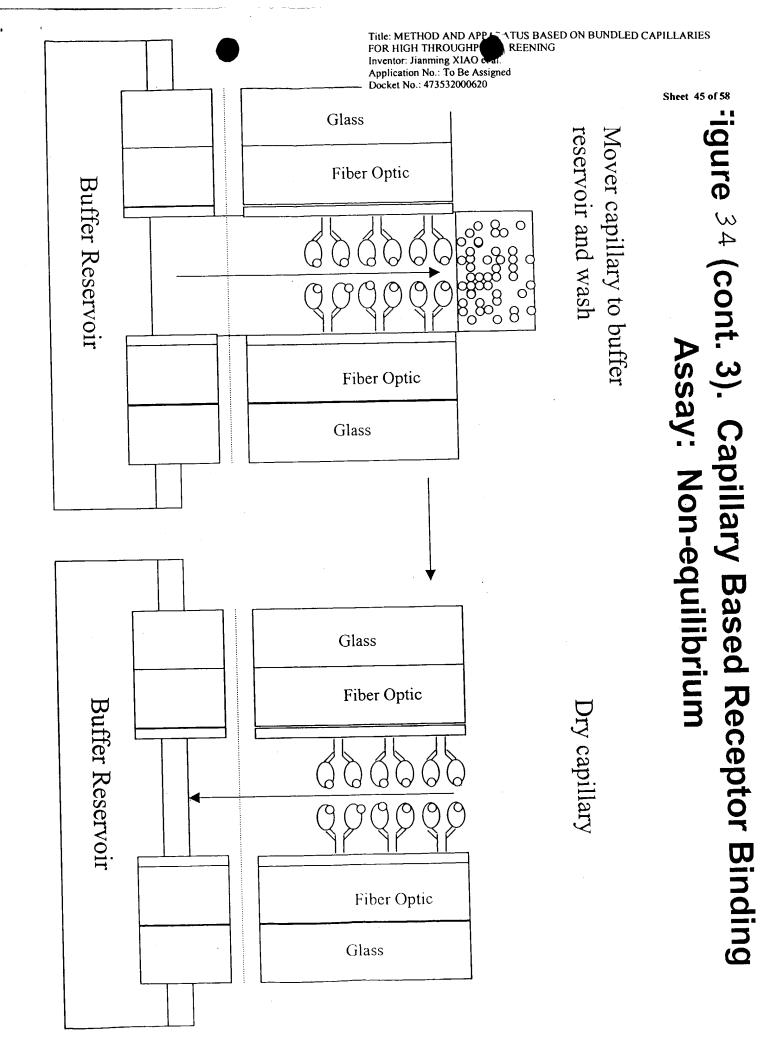
Figure 34 (cont. 2). Capillary Based Receptor Binding Assay: Non-equilibrium

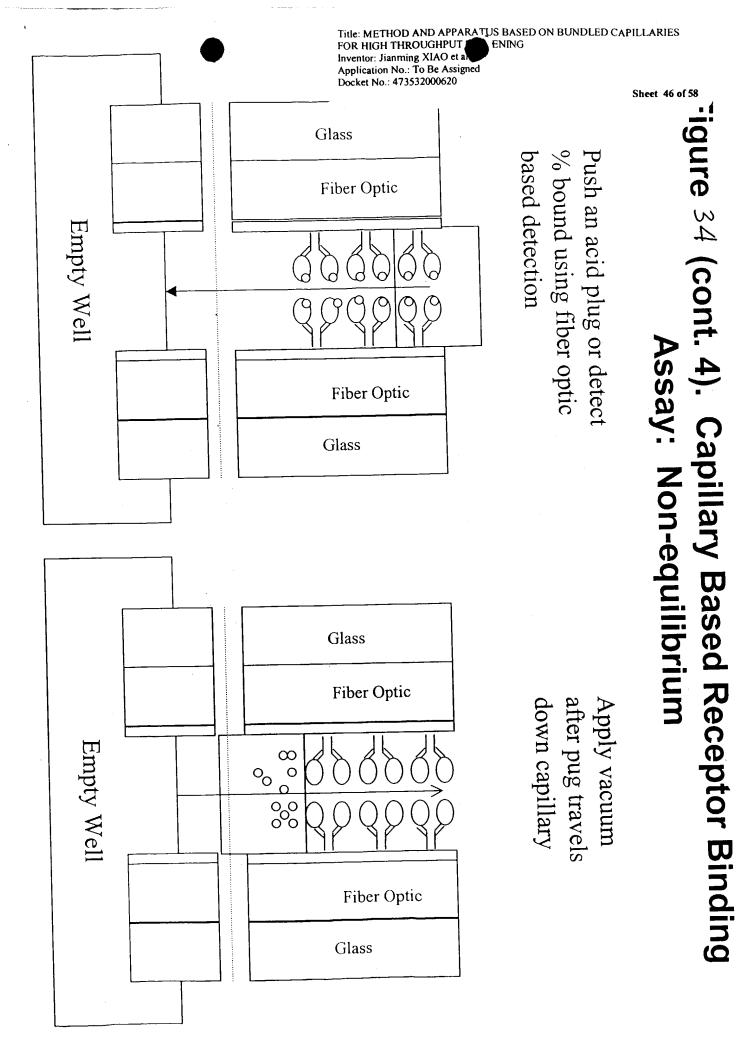
Add compound and use fiber optic based detection to observe kinetics



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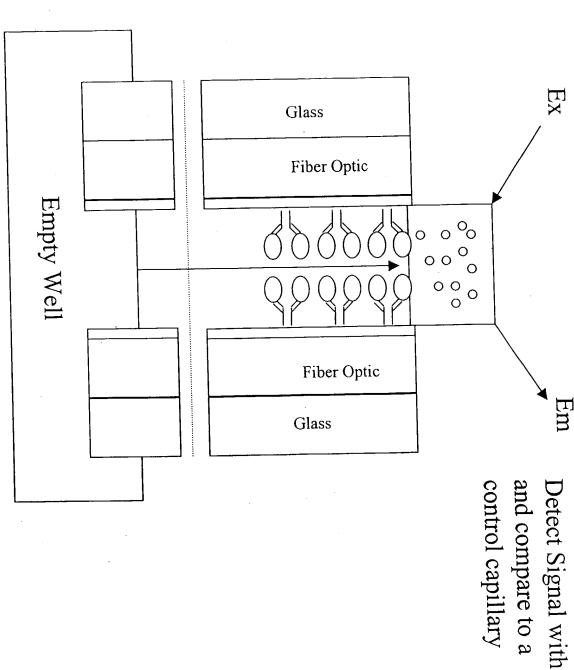


AND APPARATUS BASED ON BUNDLED CAPILLAR Title: METH FOR HIGH JUGHPUT SCREENING

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Figure 34 (cont. 5). Capillary Based Receptor Binding Assay: non-equilibrium



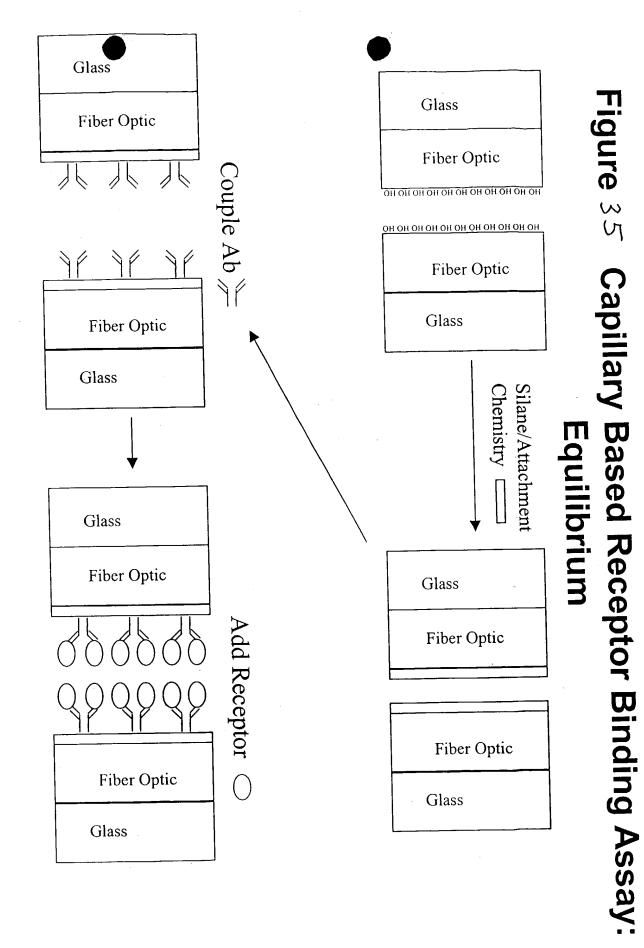
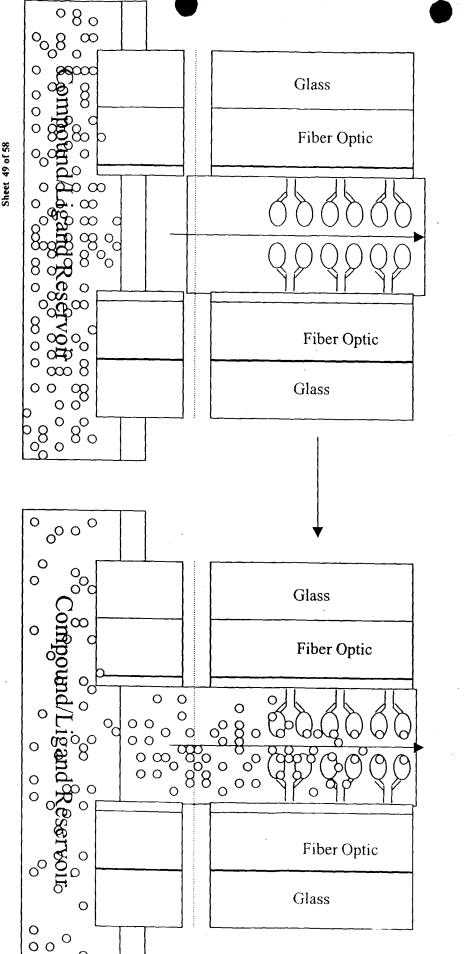


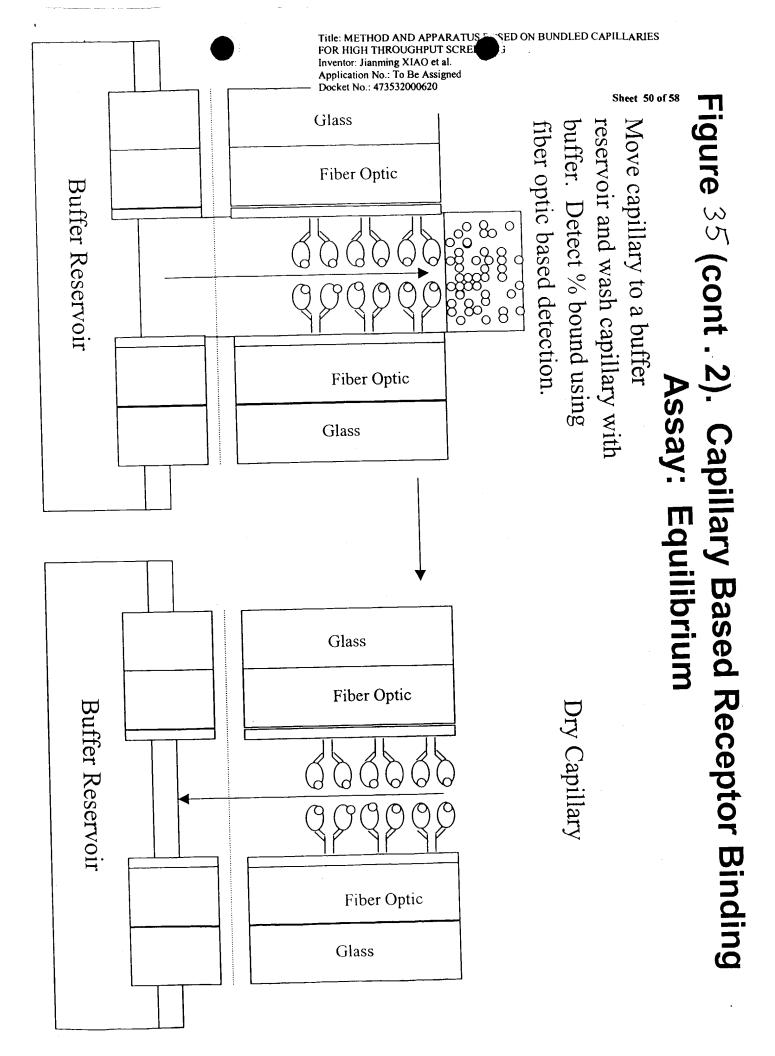
Figure 35 (cont. 1). Capillary Based Receptor Binding Assay: Equilibrium

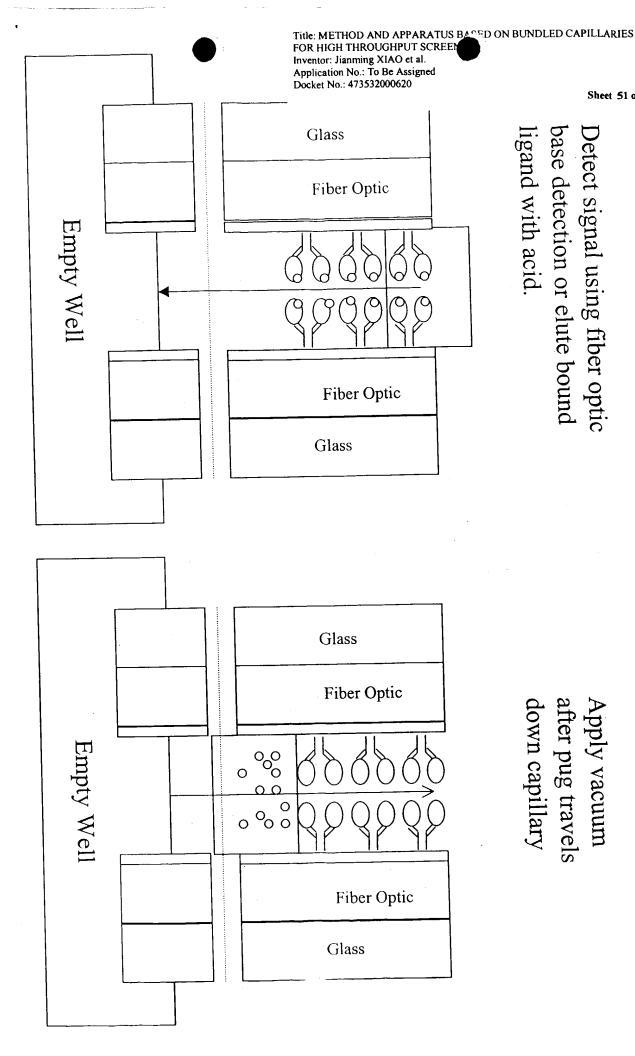
compound/ligand reservoir. Move Capillary to

using fiber optic base detection. equilibrium. Detect equilibrium Add solution and let system reach



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ligand with acid. base detection or elute bound Detect signal using fiber optic

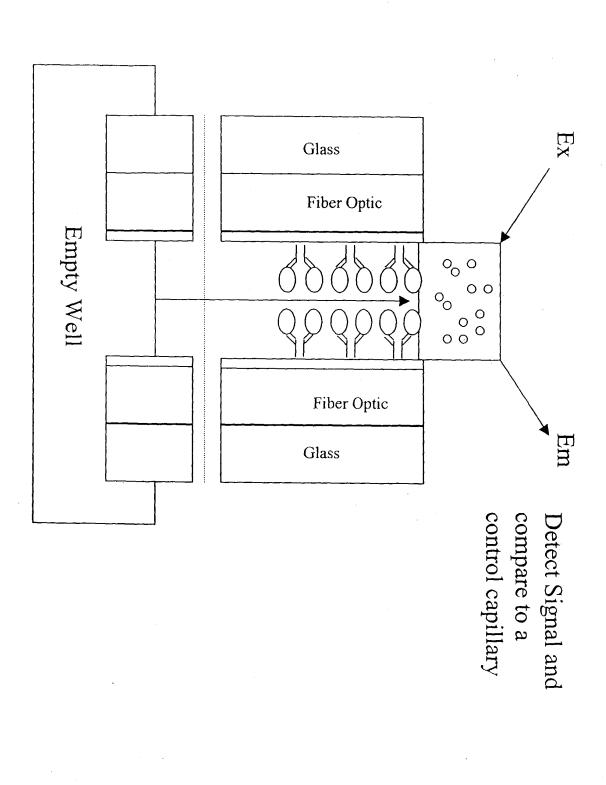
after pug travels Apply vacuum down capillary

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Title: HOD AND APPARATUS BASED ON BUNDLED CAP FOR THROUGHPUT SCREENING Inventor: Jianming XIAO et al. Application No.: To Be Assigned Docket No.: 473532000620

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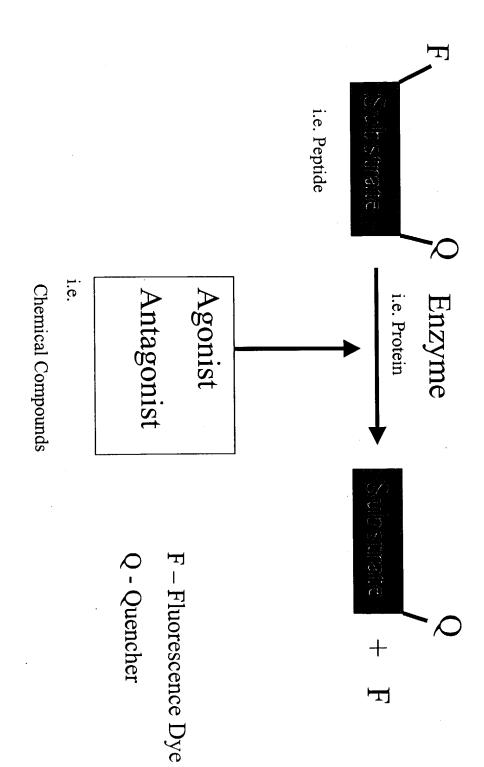
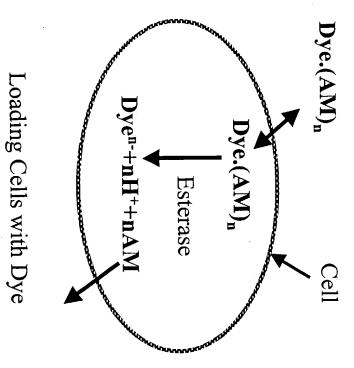


Figure 36

Title: METHOD AND APPARATUS BASED ON BUNDI TO CAPILLARIES OR HIGH THROUGHPUT SCREENING Inventor: Jianming XIAO et al.
Application No.: To Be Assigned Docket No.: 473532000620

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Agonist or

Fluorescence

Detection

Antagonist

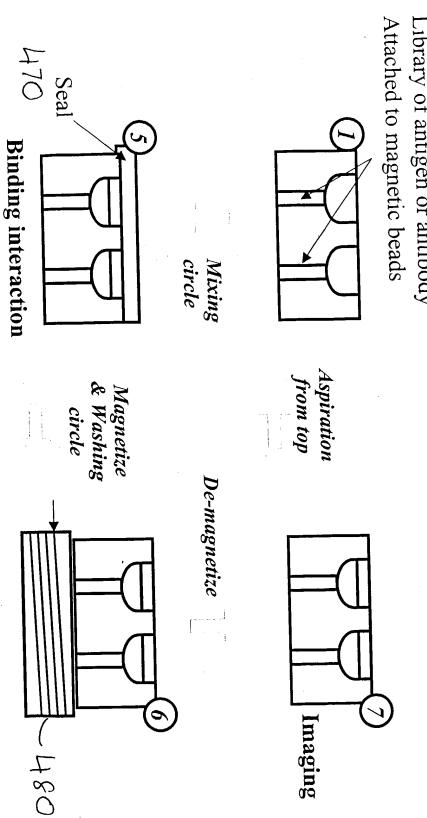
Assay Based on Tracking Cytosolic [Ca++]

Figure 37

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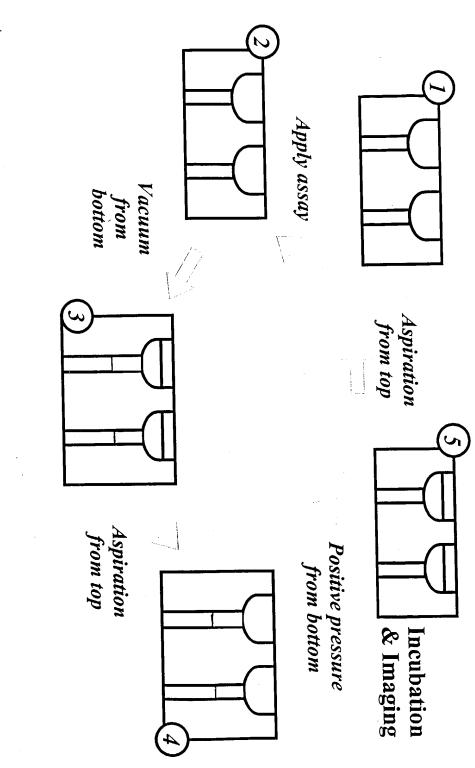
Library of antigen or antibody

Protein Array & Cell Array



Title: THOD AND APPARATUS BASED ON BUNDLED CAFOR I THROUGHPUT SCREENING
Invent.: Jianming XIAO et al.
Application No.: To Be Assigned
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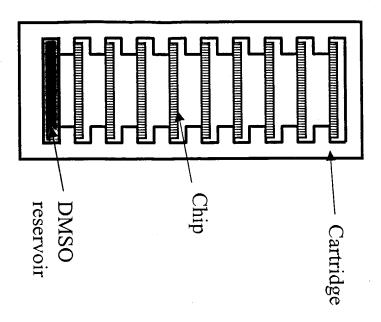


Fig. 39 One embodiment of the capillary array cartridge design

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Fig. 40 Metering with through hole plates and mixing

